

ABSTRACT

PHARMACOPHORE MODELS FOR THE IDENTIFICATION OF
THE CYP2D6 INHIBITORY POTENCY OF SELECTIVE
SEROTONIN REUPTAKE INHIBITORS

5 The present invention relates to novel screening methods which enable the
selection of selective serotonin reuptake inhibitor (SSRI) compounds which do not
possess significant inhibitory potency towards cytochrome P450 enzymes, in
particular, CYP2D6. The present invention also relates to a method of generating a
pharmacophore model for the CYP2D6 inhibitory activity of SSRI compounds; to
10 methods for the discovery of molecules that are potential SSRI compounds which do
not possess significant inhibitory potency towards the CYP2D6 enzyme; to methods
of modeling the features of the CYP2D6 pharmacophore useful in selecting SSRI's
which do not possess significant potency towards CYP2D6. Further, the invention
also relates to pharmaceutical compositions comprising an SSRI compound which
15 does not possess significant potency towards the CYP2D6 enzyme identified by
methods of the invention; to the uses of an SSRI compound identified by the methods
of the invention for the manufacture of medicaments and for the treatment of a
condition, a disorder or a disease in a mammal for which an SSRI compound
identified by the method of the invention is therapeutically useful.